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Confirmation No.: 9909 CENTRAL FAX CENTER MAY 2 1 2008

AMENDMENT TO THE CLAIMS

(Currently Amended) A method of determining the state of a disease or condition associated with endothelial cell dysfunction in a subject, the method comprising:

providing a subject suffering from or thought to be at risk of suffering from a disease or condition associated with endothelial cell dysfunction;

determining a level of profilin-1 at a site in the subject; and

relating an elevated level of profilin-1 the level to the state of the disease or condition of the subject.

- The method of claim 1, wherein an increased 2. (Original) level of profilin-1 indicates a positive diagnosis for the disease or condition.
- The method of claim 1, wherein the disease or (Original) 3. condition is diabetic vascular disease.
- The method of claim 1, wherein the disease or (Original) condition is atherosclerosis.
- The method of claim 1, wherein the disease or (Original) condition is a vascular disease selected from the cerebrovascular peripheral vascular disease, consisting of disease, cardiovascular disease and any combination thereof.
- The method of claim 1, wherein the site is (Original) associated with a vascular sample from the subject.

- 7. (Original) The method of claim 6, wherein the vascular sample is a serum sample.
- 8. (Original) The method of claim 6, wherein the vascular sample is a plasma sample.
- 9. (Original) The method of claim 1, wherein the site is associated with a tissue sample from the subject.
- 10. (Original) The method of claim 9, wherein the tissue sample from the subject is an aorta tissue sample.
- 11. (Original) The method of claim 9, wherein the tissue sample from the subject is a retina tissue sample.
- 12. (Original) A kit for performing the method of determining the state of a disease or condition associated with endothelial cell dysfunction in claim 1, the kit comprising:

an indicator, wherein the indicator is capable of indicating the increased level of profilin-1; and

- a device, wherein the device is capable of qualitatively or quantitatively monitoring the indicator for indication of the increased level of profilin-1.
- 13. (Original) A method for prophylaxis or treatment of a disease or condition associated with endothelial cell dysfunction in a subject, the method comprising:

providing a subject suffering from or thought to be at risk of suffering from a disease or condition associated with endothelial cell dysfunction;

administering to the subject a therapeutic composition or protocol to inhibit an activity or function of profilin-1,

wherein inhibiting the activity or function of profilin-1 prevents or treats the disease or condition.

- 14. (Original) The method of claim 13, wherein a pharmaceutically acceptable carrier comprises the therapeutic composition.
- 15. (Original) The method of claim 13, wherein the therapeutic protocol comprises modifying, substituting, regulating or interfering with a gene associated with profilin-1 activity or function.
- 16. (Original) The method of claim 13, wherein the therapeutic protocol comprises downregulating gene transcription associated with profilin-1 synthesis.
- 17. (Original) The method of claim 13, wherein the disease or condition is diabetic vascular disease.
- 18. (Original) The method of claim 13, wherein the disease or condition is atherosclerosis.
- 19. (Original) The method of claim 13, wherein the disease or condition is a vascular disease selected from the group consisting of cerebrovascular disease, cardiovascular disease, peripheral vascular disease and any combination thereof.
- 20. (Original) A therapeutic composition for prophylaxis or treatment of a disease or condition associated with endothelial cell dysfunction in a subject, the therapeutic composition comprising:

a inhibitor capable of inhibiting an activity or function of profilin-1; and

- a pharmaceutically acceptable carrier comprising the inhibitor.
- 21. (Original) The therapeutic composition of claim 20, wherein the disease or condition is diabetic vascular disease.
- 22. (Original) The therapeutic composition of claim 20, wherein the disease or condition is atherosclerosis.
- 23. (Original) The therapeutic composition of claim 20, wherein the disease or condition is a vascular disease selected from the group consisting of peripheral vascular disease, cerebrovascular disease and cardiovascular disease and any combination thereof.
- 24. (Original) A therapeutic protocol for prophylaxis or treatment of a disease or condition associated with endothelial cell dysfunction in a subject, the therapeutic protocol comprising:

providing a subject suffering from or thought to be atrisk of suffering from a disease or condition associated with endothelial cell dysfunction; and

administering a small interfering RNA to the subject, wherein the small interfering RNA downregulates gene transcription associated with profilin-1 to prevent or treat the disease or condition.

25. (Original) The therapeutic protocol of claim 24, wherein the small interfering RNA is a synthetic small interfering RNA.

- 26. (Original) The therapeutic protocol of claim 24 or 25, wherein the small interfering RNA is delivered via a lentivirus.
- 27. (Original) The therapeutic protocol of claim 24 or 25, wherein the small interfering RNA is delivered to an artery selected from the group consisting of coronary, carotid, femoral and a combination thereof.